Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

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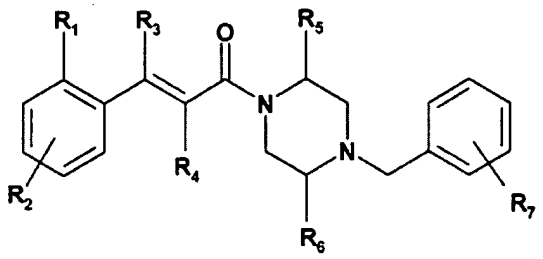
Listing of Claims:

1. (Currently Amended) A compound of formula I, or a pharmaceutically acceptable salt or ester thereof,



wherein

R_1 is -X-R_{10}, -X-$(R_{10})_2$ or –$NR_{11}R_{12}$

Wherein X is a linker ~~having 1 atom or a chain~~ having 1, 2, 3 or 4 atoms independently selected from N, C, ~~O~~ or S, and wherein when said linker has 2 or more C atoms the linker may have 1 or more C=C or C≡C bonds;

wherein any of said atoms has up to 2 optional substituents selected from hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, sulfur amino; sulfinyl, sulfonyl;

R_{10} is optionally substituted and ~~a substituent~~ is independently selected from the group consisting of hydrogen, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, cycloalkyl, heterocycloalkyl, aryl, heteroaryl;

R_{11} and R_{12} are optionally substituted and each represent a lower alkyl group connected together such that R_1 is an optionally substituted heterocycloalkyl or heteroaryl group;

R_2 and R_7 are optionally substituted and represent one or more substituents attached to the phenyl ring selected from the group consisting of hydrogen, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl, aryl, heteroaryl or a substituent forming a bicyclic ring system of which the phenyl ring to which it is attached forms part of the bicycle for example butadiene forming napthyl, or heterobutadiene forming quinolinyl, quinoxalinyl or isoquinolinyl;
R_3 and R_4 are optionally substituted and independently selected from the group consisting of hydrogen, cyano, halo, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, cycloalkyl, heterocycloalkyl, aryl;

R_5 and R_6 are optionally substituted and independently selected from the group consisting of hydrogen, cyano, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, cycloalkyl, heterocycloalkyl, aryl;

the optional substituents on X are one or more independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, amino, carbonyl, sulfur, sulfinyl, sulfonyl;

Wherein the optionally substituted substituents are optionally substituted once or more by a substituent independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro, oxy, lower alkyl, lower alkyenyl, lower alkynyl, amino, sulfur, cycloalkyl, heterocyloalkyl, aryl;

the optional substituents on R_{10} are one or more substituents independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl, aryl; wherein the optionally substituted substituents are optionally substituted once or more by a substituent independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl, aryl;

the optional substituents on R_{11} and R_{12} are one or more substituents independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl, aryl;

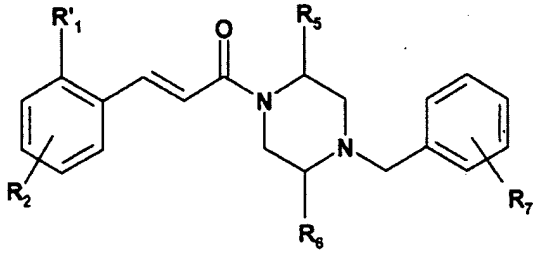
the optional substituents on R_2 and R_7 are one or more substituents independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl, aryl;

wherein the optionally substituted substituents are optionally substituted once or more by a substituent independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl, aryl;

the optional substituents on R_3 and R_4 are one or more substituents independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl, aryl;
wherein the optionally substituted substituents are optionally substituted once or more by a substituent independently selected from the group consisting of hydrogen, oxo, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl, aryl;

the optional substituents on R_5 and R_6 are one or more substituents independently selected from the group consisting of hydrogen, oxo, cyano, hydroxyl, optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, cycloalkyl, heterocycloalkyl, aryl, imino, oxime;
wherein the optionally substituted substituents are optionally substituted once or more by a substituent independently selected from the group consisting of hydrogen, oxo, hydroxyl, cyano, halo, nitro or optionally substituted oxy, lower alkyl, lower alkyenyl, lower alkynyl, carbonyl, amino, sulfur, cycloalkyl, heterocycloalkyl~~,~~ or aryl~~;~~
~~with the proviso that compounds wherein X is -O-C-C-C- are excluded~~.

2. (Previously Presented) A compound of formula II, or a pharmaceutically acceptable salt or ester thereof,

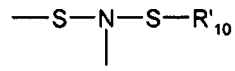


II

wherein

R'_1 is $-X'-R'_{10}$

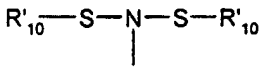
Wherein X' is a linker independently selected from optionally substituted –N-C-N-, -N-C-, -N-S-, -N-S-N-, -C-N-, -S-N-, -C≡C-, -C=C-, -N-C-S-, -C-, or



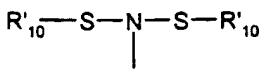
wherein $R_2 - R_{10}$ are as herein before defined;

R'_{10} is one or more substituents independently selected from the group consisting of hydrogen, halo, optionally substituted methyl, optionally substituted isopropyl, optionally substituted imidazolyl or thiazolyl, 3-oxa-1-aza-spiro[4.4]nonan-2-one, hydroxy, optionally substituted pyrrolidinyl, morpholino, piperazinyl, formic acid methyl ester, [1,2,4]triazol, imidazolidinyl, tetrazolyl, $-N(CH_3)-OCH_3$ or methoxy, optionally substituted carbonyl, amino, heterocycloalkyl and aryl;

when R'_1 is $-N-C-N-R'_{10}$ the C atom is substituted by oxo, =N-C≡N or $=C-NO_2$;
when R'_1 is $-N-C-N-R'_{10}$, R'_{10} is hydrogen;
when R'_1 is $-N-C-N-R'_{10}$, R'_{10} is optionally substituted by hydrogen;
when R'_1 is $-N-C-R'_{10}$ or $-C-N-R'_{10}$ the C atom is substituted by oxo;
when R'_1 is $-N-C-R'_{10}$ or $-C-N-R'_{10}$, R'_{10} is optionally substituted methyl, piperidinyl, imidazolidinyl, pyrrolidinyl, morpholino;
when R'_1 is $-N-C-R'_{10}$ or $-C-N-R'_{10}$, R'_{10} is substituted by hydrogen, methyl, benzyl, acetyl, oxo, dimethylamino, isopropyl, hydroxy, formic acid ethyl ester;
when R'_1 is $-N-S-R'_{10}$ or

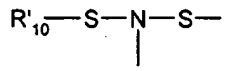


the S atom or atoms are substituted twice by oxo;
when R'_1 is $-N-S-R'_{10}$ or



R'_{10} is optionally substituted methyl;

when R'_1 is -N-S-R'_{10} or



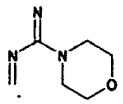
R'_{10} is optionally substituted by hydrogen;

when R'_1 is -N-S-N-R'_{10} the S atom is substituted twice by oxo and the N atom is independently optionally substituted by methyl;
when R'_1 is -N-S-N-R'_{10}, R'_{10} is hydrogen or optionally substituted methyl, imidazolyl, thiazolyl;
when R'_1 is -N-S-N-R'_{10}, R'_{10} is optionally substituted by hydrogen, methyl, acetamidyl;

when R'_1 is -C≡C-R'_{10}, R'_{10} is optionally substituted methyl, isopropyl or piperindinyl;
when R'_1 is -C≡C-R'_{10}, R'_{10} is optionally substituted by hydrogen or amine;

when R'_1 is -C=C-R'_{10}, R'_{10} is optionally substituted piperidinyl,
when R'_1 is -C=C-R'_{10}, R'_{10} is optionally substituted by hydroxy, methyl;

when R'_1 is -N-C-S-R'_{10} the C atom is substituted by =N-C≡N or

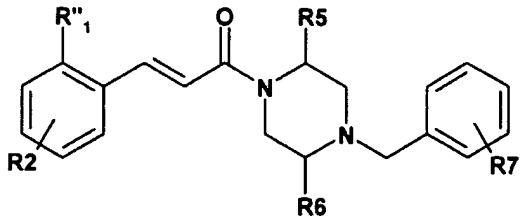


when R'_1 -N-C-S-R'_{10}, R'_{10} is optionally substituted methyl,
when R'_1 is -N-C-S-R'_{10}, R'_{10} is optionally substituted by hydrogen;

when R'_1 is -C-R'_{10} the C atom is optionally substituted by oxo,
when R'_1 -C-R'_{10}, R'_{10} is 3-oxa-1-aza-spiro[4.4]nonan-2-one, hydroxy, optionally substituted pyrrolidinyl, morpholino, piperazinyl, formic acid methyl ester, [1,2,4]triazol, imidazolidinyl, tetrazolyl, -N(CH_3)-OCH_3 or methoxy;
when R'_1 is -C-R'_{10}, R'_{10} is optionally substituted by hydrogen, oxo, methyl, acetyl, isopropyl, methoxy, hydroxy, formic acid methyl ester, dimethylamino or ethanone;

The optional substituents on R'_{10} are one or more substituents independently selected from the group consisting of hydrogen, or optionally substituted oxy, lower alkyl, carbonyl, amino; Wherein the optionally substituted substituents are optionally substituted once or more by a substituent independently selected from the group consisting of hydrogen, optionally substituted oxy; or optionally substituted lower alkyl.

3. (Original) A compound of formula IIa, or a pharmaceutically acceptable salt or ester thereof,



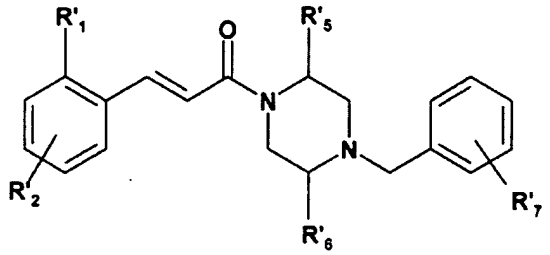
IIa

wherein

$R"_1$ is $-NR"_{11}R"_{12}$

Wherein $-NR"_{11}R"_{12}$ collectively represents imidazolidinyl-2,4-dione, optionally substituted once or twice by a lower alkyl group.

4. (Previously Presented) A compound of formula III, or a pharmaceutically acceptable salt or ester thereof,



III

wherein R'_1 is as herein before defined,

R'_2 and R'_7 are hydrogen, cyano, halo, butadienyl, methoxy, ethoxy, 2-methoxyethoxy, morpholino, trifluoromethoxy, 2-methylpropoxy, 2-propoxy;

R'_5 and R'_6 are independently selected from the group consisting of hydrogen and lower alkyl, acetyl.

5.-8. (Canceled)

9. (Original) A pharmaceutical composition comprising a compound according to claim 1 in association with a pharmaceutically acceptable diluent or carrier.

10. (Previously Presented) A compound of claim 1 selected from the group consisting of:
N-(5-Chloro-2-[(E)-3-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]phenyl)-N'-cyanoguanidine,
N-(5-Chloro-2-[(E)-3-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]phenyl)-acetamide,

N-(5-Chloro-2-[(E)-3-[(S)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]phenyl)-acetamide,

(5-Chloro-2-[(E)-3-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-urea,

(5-Chloro-2-[(E)-3-[(S)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-urea,

N-(5-Chloro-2-[(E)-3-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]phenyl)-N,N-dimethylsulfamide,

N-(5-Chloro-2-[(E)-3-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]phenyl)-methanesulfonamide,

1-Acetyl-piperidine-4-carboxylic acid (5-chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-amide,

1- Methyl-1H–imidazole-4-sulfonic acid(5-Chloro-2-[(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl]-phenyl)-amide,

N-[5-(5-Chloro-2-[(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl]-phenylsulfamoyl)-thiazol-2yl]-acetamide,

2-Oxo-imidazolidine-1-carboxylic acid (5-Chloro-2-[(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl]-phenyl)-amide,

N-(4-Chloro-2-[(E)-3-[4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-methylthio-N'-cyano thiourea,

N-(4-Chloro-2-[(E)-3-[4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-sulfonylurea,

(5-Chloro-2-[(E)-3-[(2R,5S)-4-(4-fluorobenzyl)-2,5-dimethylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-urea,

(5-Chloro-2-[(E)-3-[(2R,5S)-4-(4-fluorobenzyl)-2,5-dimethylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-urea,

N-(5-Chloro-2-[(E)-3-[(2R,5S)-4-(4-fluorobenzyl)-2,5-dimethyl-piperazin-1-yl]-3-oxopropenyl]-phenyl)- acetamide,

N-(5-Chloro-2-[(E)-3-[(2S,5R)-4-(4-fluorobenzyl)-2,5-dimethyl-piperazin-1-yl]-3-oxopropenyl]-phenyl)- acetamide,

(5-Chloro-2-[(E)-3-[(2S,5R)-4-(4-fluorobenzyl)-2,5-dimethylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-urea,

N-(5-Chloro-2-[(E)-3-[-4-(4-fluorobenzyl)-piperazin-1-yl]-3-oxopropenyl]-phenyl)-acetamide,

(5-Chloro-2-[(E)-3-[-4-(4-fluorobenzyl)-piperazin-1-yl]-3-oxopropenyl]-phenyl)-urea,

(E)-3-[4-Chloro-2-(4-hydroxy-1-methylpiperidin-4-ylethynyl)-phenyl]-1-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-propenone,

(E)-3-[4-Chloro-2-(4-hydroxy-1-methylpiperidin-4-ylethynyl)-phenyl]-1-[(S)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-propenone,

(E)-3-[4-Chloro-2-[(E)-2-(4-hydroxy-1-methylpiperidin-4-yl)-vinyl]-phenyl]-1-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-propenone,

(E)-3-[4-Chloro-2-[(E)-2-(4-hydroxy-1-methylpiperidin-4-yl)-vinyl]-phenyl]-1-[(S)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-propenone,

4-(5-Chloro-2-[(E)-3-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-phenylethynyl)-4-hydroxypiperidine-1-carboxylic acid tert butyl ester,

(E)-3-[4-Chloro-2-(4-hydroxypiperidin-4-ylethynyl)-phenyl]-1-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-propenone,

(E)-3-[2-(3-Amino-3-methylbut-1-ynyl-4-chlorophenyl]-1-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-propenone,

(E)-3-[4-Chloro-2-(3-dimethylaminoprop-1-ynyl)-phenyl]-1-[4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-propenone,

(E)-3-[4-Chloro-2-(3-hydroxy-3-methylbut-1-ynyl)-phenyl]-1-[4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-propenone,

N-(3-[(E)-3-[(R)-4-(4-Fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-naphthalen-2-yl)-acetamide,

(3-[(E)-3-[(R)-4-(4-Fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-naphthalen-2-yl)-urea,

N-(3-[(E)-3-[(R)-4-(4-Fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-naphthalen-2-yl)-N'-cyanoguanidine,

N-(4-Chloro-2-[(E)-3-[4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-N'-cyanoguanidine,

N-(4-Chloro-2-[(E)-3-[4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-acetamide,

N-(6-[(E)-3-[(R)-4-(4-Fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-quinolin-7-yl)-acetamide,

(6-[(E)-3-[(R)-4-(4-Fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-quinolin-7-yl)-urea,

N-(7-[(E)-3-[(R)-4-(4-Fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-quinolin-6-yl)-acetamide,

2-Dimethylamino-N-(7-[(E)-3-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-quinolin-6-yl)-acetamide,

N-(7-[(E)-3-[(R)-4-(4-Fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-quinolin-6-yl)-methanesulfonamide,

N-(5-Chloro-2-[(E)-3-[(2R,5S)-4-(4-fluorobenzyl)-2,5-dimethylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-cyanoguanidine,

N-(5-Chloro-2-[(E)-3-[(2R,5S)-4-(4-fluorobenzyl)-2,5-dimethylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-2-dimethylacetamide,

N-(5-Chloro-2-[(E)-3-[(2R,5S)-4-(4-fluorobenzyl)-2,5-dimethylpiperazin-1-yl]-3-oxopropenyl]-phenyl)-methanesulfonamide,

N-(5-Chloro-2-[(E)-3-[(2R,5S)-4-(4-fluorobenzyl)-2,5-dimethylpiperazin-1-yl]-3-oxopropenyl]-4-methoxyphenyl)-acetamide,

N-(5-Chloro-2-[(E)-3-[(2R,5S)-4-(4-fluorobenzyl)-2,5-dimethylpiperazin-1-yl]-3-oxopropenyl]-4-methoxyphenyl)-methanesulfonamide,

N-[5-Chloro-2-[(E)-3-(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl)-4-(2-methoxyethoxy)-phenyl]-acetamide,

N-(5-Chloro-2-[(E)-3-[(R)-4-(4-fluorobenzyl)-2-methylpiperazin-1-yl]-3-oxopropenyl]-4-morpholin-4-yl-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[(R)-2-ethyl-4-(4-fluorobenzyl)-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

(5-Chloro-2-{(E)-3-[(R)-2-ethyl-4-(4-fluoro-benzyl)-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-urea,

N-(5-Chloro-4-ethoxy-2-{(E)-3-[(R)-4-(4-fluorobenzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

(5-Chloro-4-ethoxy-2-{(E)-3-[(R)-4-(4-fluorobenzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-urea,

N-(5-Chloro-4-ethoxy-2-{(E)-3-[(R)-4-(4-fluorobenzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-methanesulfonamide,

5-Oxo-pyrrolidine-2-carboxylic acid (5-chloro-4-ethoxy-2-{(E)-3-[®-4-(4-fluorobenzyl)-2-methyl-piperazin-1yl]-3oxo-propenyl}-phenyl)-amide,

N-(5-Chloro-2-{(E)-3-[(S)-4-(4-fluoro-benzyl)-2-hydroxymethyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[(S)-4-(4-fluoro-benzyl)-2-hydroxymethyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-methanesulfonamide,

(5-Chloro-2-{(E)-3-[(S)-4-(4-fluoro-benzyl)-2-hydroxymethyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-urea,

5-Oxo-pyrrolidine-2-carboxylic acid (5-chloro-2-{(E)-3-[(S)-4-(4-fluoro-benzyl)-2-hydroxymethyl-piperazin-1-yl]-3-propenyl}-4-methoxy-phenyl)-amide,

N-(5-chloro-2-{(E)-3-[(S)-4-(4-fluoro-benzyl)-2-hydorxymethyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)acetamide,

N-(2-{(E)-3-[(R)-2-Aminomethyl-4-(4-fluoro-benzyl)-piperazin-1-yl]-3-oxo-propenyl}-5-chloro-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[(S)-4-(4-fluoro-benzyl)-2-((S)-1-hydroxy-ethyl)-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

N-(2-{(E)-3-[(S)-2-Acetyl-4-(4-fluoro-benzyl)-piperazin-1-yl]-3-oxo-propenyl}-5-chloro-phenyl)-acetamide,

N-{5-Chloro-2-[(E)-3-((S)-4-(4-fluoro-benzyl)-2-{1-[hydroxyimino]-ethyl}-piperazin-1-yl)-3-oxo-propenyl}-acetamide,

N-(2-{(E)-3-[(2S,5S)-2-Benzyloxymethyl-4-(4-fluoro-benzyl)-5-methyl-piperazin-1-yl]-3-oxo-propenyl}-5-chloro-phenyl)-acetamide,

(S)-1-Acetyl-pyrrolidine-2-carboxylic acid (5-chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-amide,

(S)-1-Isopropyl-pyrrolidine-2-carboxylic acid (5-chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-amide,

(R)-1-Isopropyl-pyrrolidine-2-carboxylic acid (5-chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-amide,

(2S,4R)-1-Acetyl-4-hydroxy-pyrrolidine-2-carboxylic acid (5-chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-amide,

(E)-3-(4-Chloro-2-morpholin-4-ylmethyl-phenyl)-1-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

1-(5-Chloro-2-{(E)-3-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzyl)-pyrrolidin-2-one,

(E)-3-(4-Chloro-2-[1,2,4]triazol-1-ylmethyl-phenyl)-1-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

(E)-3-[4-Chloro-2-(4-methyl-piperazin-1-ylmethyl)-phenyl]-1-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

(E)-3-[2-(4-Acetyl-piperazin-1-ylmethyl)-4-chloro-phenyl]-1-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

(E)-3-[4-Chloro-2-(4-isopropyl-piperazin-1-ylmethyl)-phenyl]-1-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

1-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzyl)-3-oxa-1-aza-spiro[4.4]nonan-2-one,

3-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzyl)-5,5-dimethyl-imidazolidine-2,4-dione,

3-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzyl)-1-methyl-imidazolidine-2,4-dione,

(E)-3-[4-Chloro-2-(5-methyl-tetrazol-1-ylmethyl)-phenyl]-1-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-N-methoxy-N-methyl-benzamide,

5-Chloro-2-{(E)-3-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzoic acid methyl ester,

(5-Chloro-2-{(E)-3-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetic acid methyl ester,

5-Chloro-2-{(E)-3-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzoic acid,

5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzoic acid,

(E)-3-[4-Chloro-2-(4-methyl-piperazine-1-carbonyl)-phenyl]-1-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

5-Chloro-2-{(E)-3-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-N-isopropyl-benzamide,

5-Chloro-2-{(E)-3-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-N-(1-methyl-piperidin-4-yl)-benzamide,

N-(1-Benzyl-piperidin-4-yl)-5-chloro-2-{(E)-3-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzamide,

4-(5-Chloro-2-{(E)-3-[4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzoylamino)-piperidine-1-carboxylic acid ethyl ester,

(2S,4R)-1-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-benzoyl)-4-hydroxy-pyrrolidine-2-carboxylic acid methyl ester,

(E)-3-[4-Chloro-2-((R)-3-dimethylamino-pyrrolidine-1-carbonyl)-phenyl]-1-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

(E)-3-[4-Chloro-2-((S)-3-dimethylamino-pyrrolidine-1-carbonyl)-phenyl]-1-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

(E)-3-[2-(4-Acetyl-piperazine-1-carbonyl)-4-chloro-phenyl]-1-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

N-(5-Chloro-2-{(E)-3-[4-(4-chloro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[4-(3-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[4-(2,4-difluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[4-(4-cyano-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-prop-enyl}-4-methoxy-phenyl)-acetamide,

N-(5-Chloro-4-fluoro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-urea,

N-(5-Chloro-4-fluoro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-methanesulfonamide,

(5-Chloro-4-fluoro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-urea,

N-(5-Chloro-4-cyano-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-trifluoromethoxy-phenyl)-acetamide,

(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-trifluoromethoxy-phenyl)-urea,

N-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-isobutoxy-phenyl)-acetamide,

N-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-isopropoxy-phenyl)-acetamide,

3-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-5,5-dimethyl-imidazolidine-2,4-dione,

3-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-imidazolidine-2,4-dione,

3-(5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-1,3-diaza-spiro[4.4]nonane-2,4-dione,

3-(5-Chloro-4-fluoro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-1,3-diaza-spiro[4.5]decane-2,4-dione,

3-(5-Chloro-4-fluoro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-phenyl)-5,5-dimethyl-imidazolidine-2,4-dione,

Morpholine-4-carboxylic acid (5-chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-amide,

Pyrrolidine-1-carboxylic acid (5-chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-phenyl)-amide,

5-Chloro-2-{(E)-3-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-3-oxo-propenyl}-4-methoxy-benzoic acid methyl ester, and

(E)-3-[2-(4-Acetyl-piperazine-1-carbonyl)-4-chloro-5-methoxy-phenyl]-1-[(R)-4-(4-fluoro-benzyl)-2-methyl-piperazin-1-yl]-propenone,

or a pharmaceutically acceptable salt, or ester thereof.